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MDTKHFLPLDFSTQVNSSLTSPTGRGSMAAPSLHPSLGP GIGSPGQLHSPISTLSSPINGMG
PPFSVISSPMGPHSMSVPTTPTLGFSTGSPQLSSPMNPVSSSEDIKPPLGLNGVLKVPAPHS
GNMASFTKHICAICGDRSSGKH YGVYSCEGCKGFFKRTVRKDLTYTCRDNKDCLIDKRQRNR
CQYCRYQKCLAMGMKREAVQEERQRGKDRNENEVESTSSANEDMPVERILEAELAVEPKTET
YVEANMGLNPSSPNDPVTNICQAADKQLFTLVEWAKRIPHFSELPLDDQVILLRAGWNELLI
ASFHSRISIAVKDGILLATGLHVHRNSAHSAGVGAI FDRVLTEL VSKMRDMQMDKTEL GCLRA
IVLFNPDSKGLSNPAEVEALREKVYASLEAYCKHKYPEQPGRFAKLLLRLPALRSIGLKCLE
HLFFFKLIGDTPIDTFLMEMLEAPHQMT

FIGURE 1A

MSWAARPPFLPQRHAAGQCGPVGVRKEMHCGVASRWRRRRPWLDPAAAAA AVAGGEQQTPE
PEPGEAGRDGMGDSGRDSRSPDSSSPNLPQGVPPPSPPGPPLPPSTAPSLGGSGAPPPPPM
PPPPLGSPFPVISSSMGSPGLPPPAPPGFSGPVSSPQINSTVSLPGGGSGPPEDVKPPVLGV
RGLHCPPPPGGPGAGKRLCAICGDRSSGKH YGVYSCEGCKGFFKRTIRKDLTYSCRDNKDCT
VDKRQRNRCQYCRYQKCLATGMKREAVQEERQRGKDKDGDGEGAGGAPEEMPVDRILEAELA
VEQKSDQGVVEGPGGTGGSGSSPNDPVTNICQAADKQLFTLVEWAKRIPHFSSPLDDQVILL
RAGWNELLIASFHSRISIDVRDGILLATGLHVHRNSAHSAGVGAI FDRVLTEL VSKMRDMRMD
KTEL GCLRAIILFNPD AKGLSNPSEVEVLREKVYASLE TYCKQKYPEQQGRFAKLLLRLPAL
RSIGLKCLEHLFFFKLIGDTPIDTFLMEMLEAPHQLA

FIGURE 1B

MYGNYSHFMKFPAGYGGSPGHTGSTSMSPSAALSTGKPMDSHPSYTDTPVSAPRTL SAVGTP
LNALGSPYRVITSAMGPPSGALAAPPGINLVAPPSSQLNVVNSVSSSEDIKPLPGLPGIGNM
NYPSTSPGSLVKHICAICGDRSSGKH YGVYSCEGCKGFFKRTIRKDLIYTCRDNKDCLIDKR
QRNRCQYCRYQKCLVMGMKREAVQEERQSRERA ESEAECATSGHEDMPVERILEAELAVEP
KTESYGD MNMENSTNDPVTNICHAADKQLFTLVEWAKRIPHFSDLTLEDQVILLRAGWNELL
IASFSHRSVSVQDGILLATGLHVHRSSAHSAGVGSIFDRVLTEL VSKMKDMQMDKSELGCLR
AIVLFNPDAKGLSNPSEVETLREKVYATLEAYTKQKYPEQPGRFAKLLLRLPALRSIGLKCL
EHLFFFKLIGDTPIDTFLMEMLETPLQIT

FIGURE 1C

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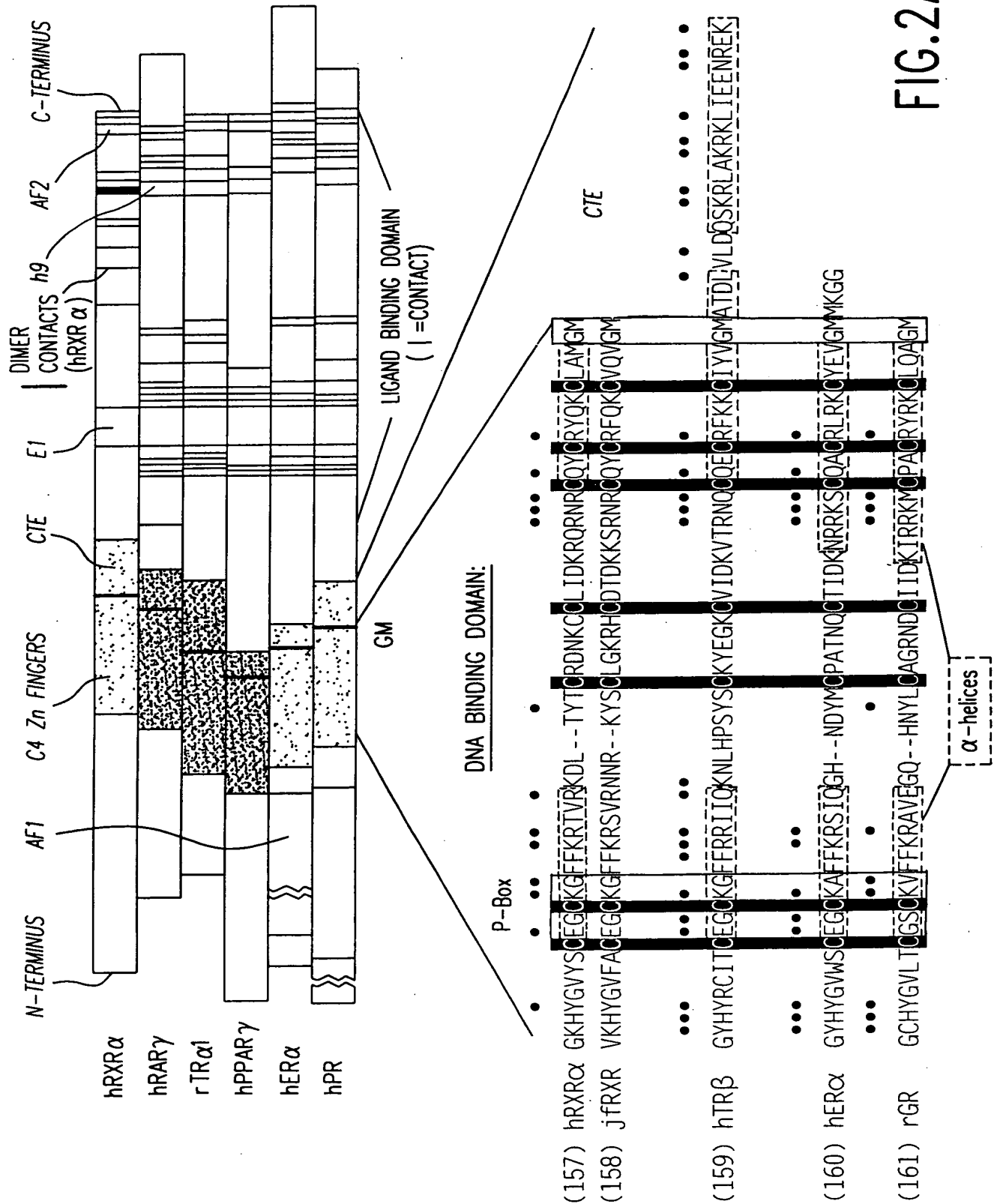


FIG. 2A

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E1 Region		Heptad 9 (h9)		AF2 Domain	
		ϕ	ϕ	ϕ	ϕ
(162) hRXR α	WAKRIPHFSELPLDQVILL	(172) hRXR α	LLRLPALR	(182) hRXR α	FLMEML
(163) jFRXR	WAKRLPHFRDLSIADQVLL	(173) jFRXR	VILRIPALR	(183) jFRXR	FLLDML
(164) hTR α 1	FAKKLPMFSELPCEQIILL	(174) hTR α 1	LLMKVTDLR	(184) hTR α 1	LFLEVF
(165) hRAR γ	FAKRLPGFTGLSIADQITLL	(175) hRAR γ	MLMKITDLR	(185) hRAR γ	LIREML
(166) hPPAR γ	YAKSIPGFVNLDLNDQVTLL	(176) hPPAR γ	LLQKMTDLR	(186) hPPAR γ	LLOEIV
(167) hLXR	FAKQLPGFLQLSREDQIALL	(177) hLXR	MLMKLVSLR	(187) hLXR	LLSEIW
(168) hVDR	FAKMIPGFRDLTSEDQIVLL	(178) hVDR	MIQKLADLR	(188) hVDR	LVLEVF
(169) hER	WAKRVPGFVDLTLDQVHLL	(179) hER	LLLILSHIR	(189) hER	LLLEML
(170) hGR	WAKAIPGFRNLHLDQDMTLL	(180) hGR	LTKLLDSMH	(190) hGR	MLAEII
(171) hPR	WSKSLPGFRNLHIDDQITLI	(181) hPR	LTKLLDNLH	(191) hPR	MMSEVI
α -helix 3		α -helix 10/11		α -helix 12	

FIG.2B

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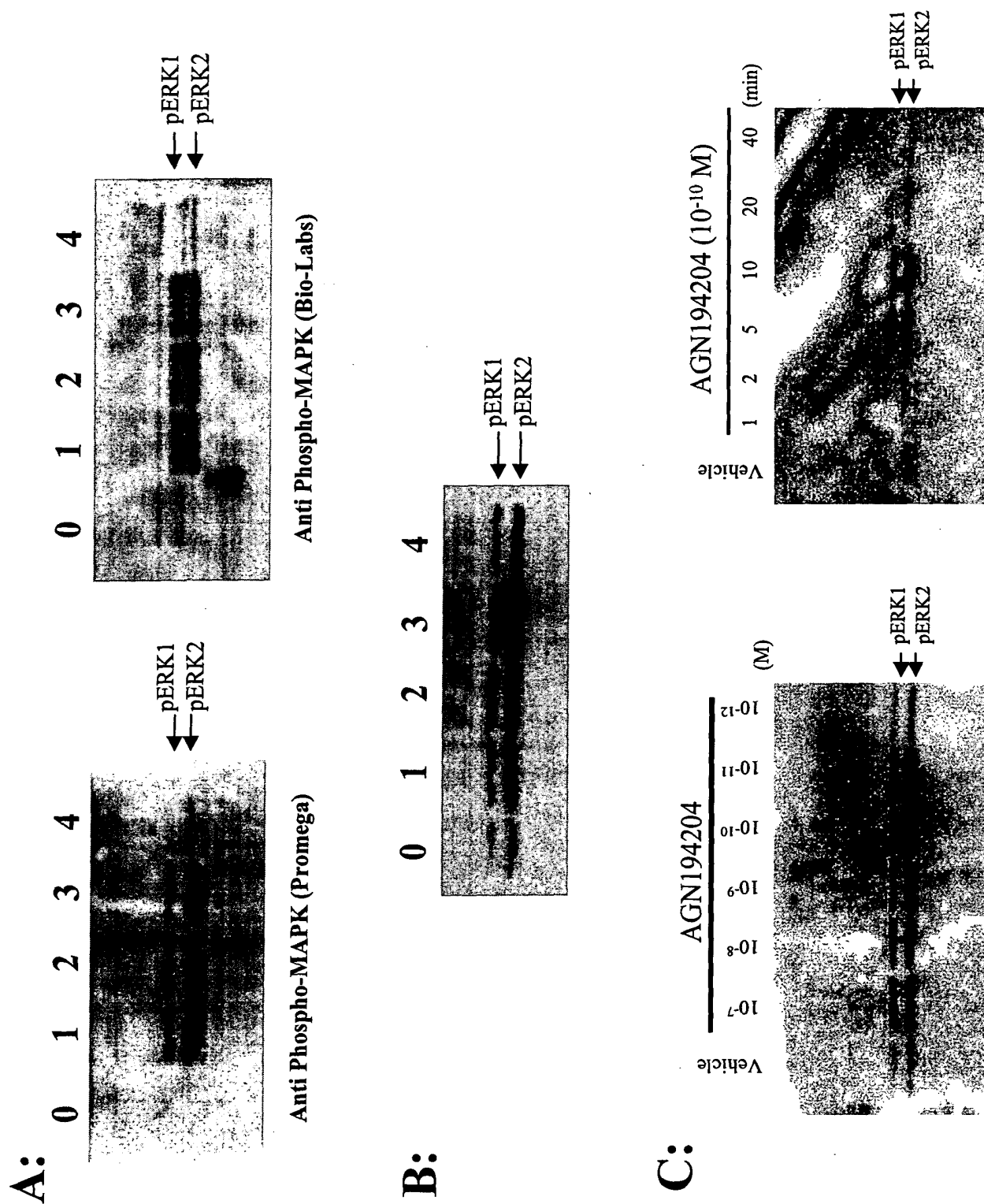


FIGURE 3

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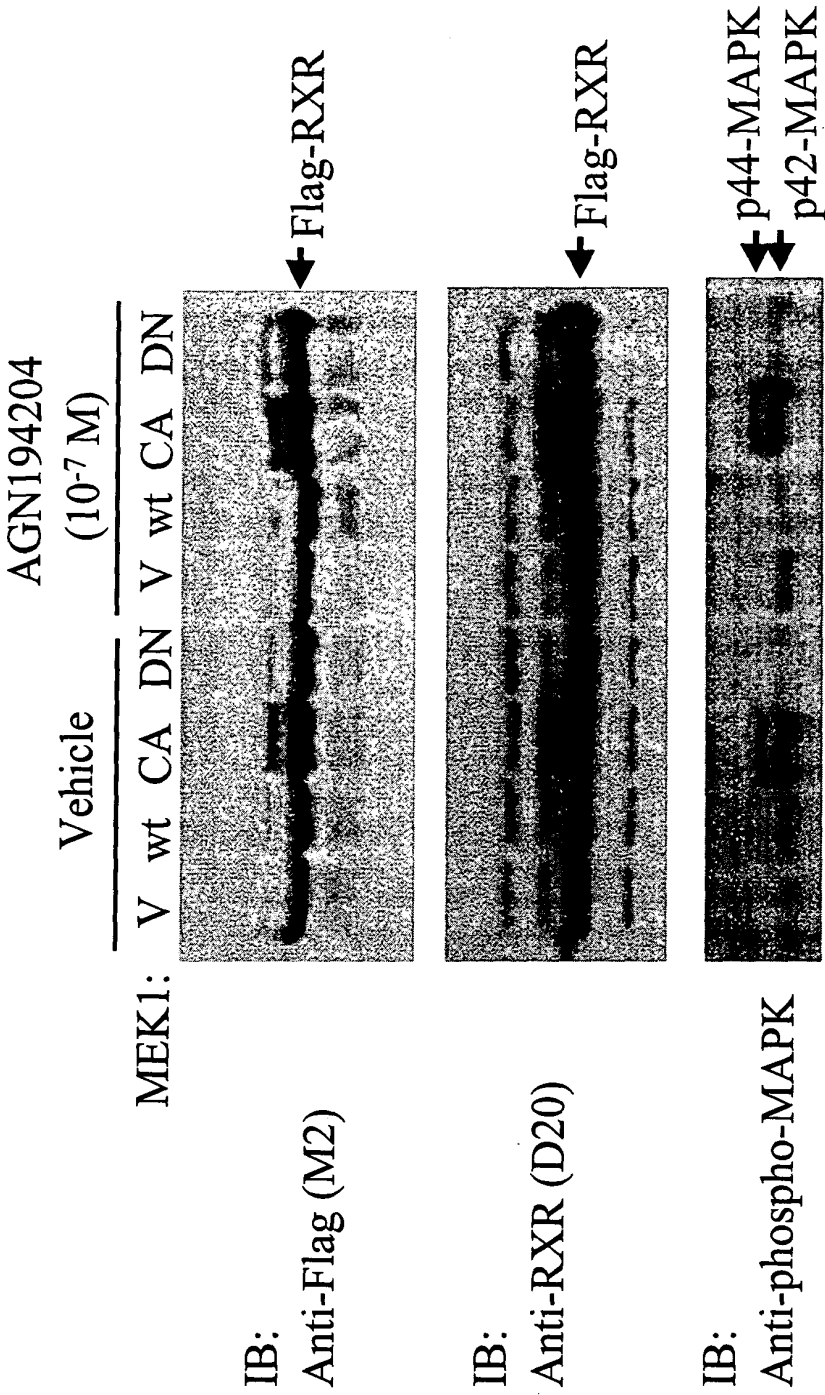


FIGURE 4

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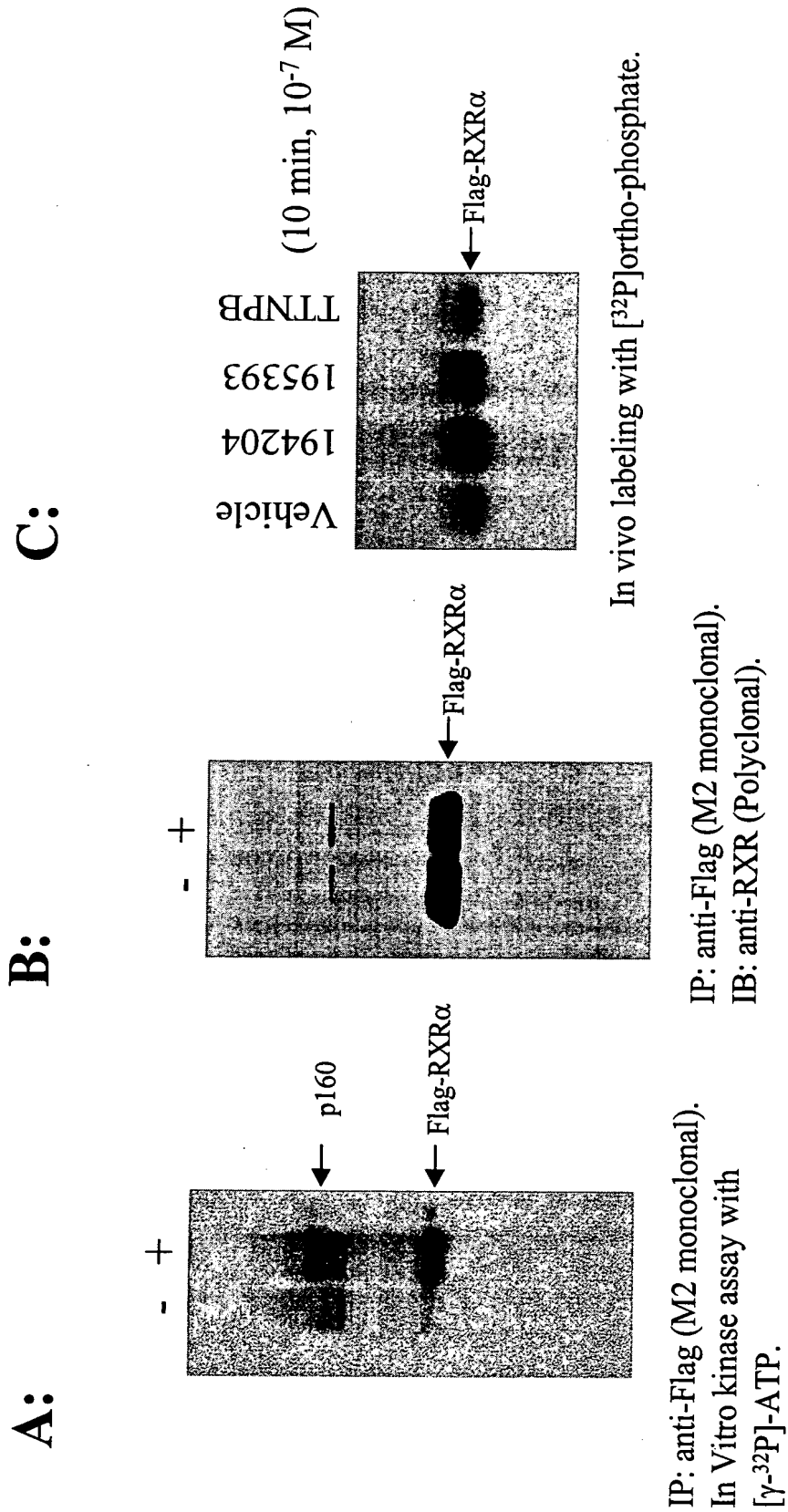
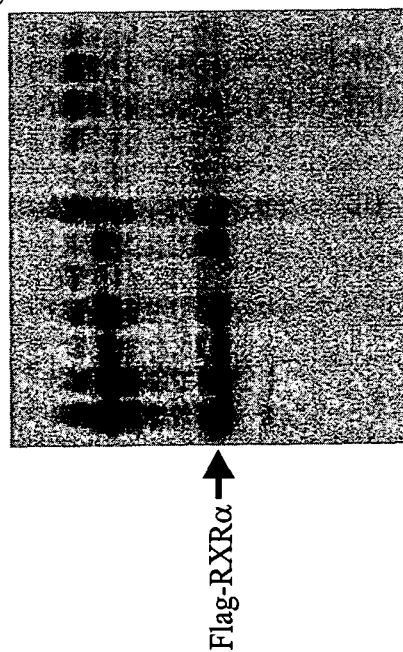


FIGURE 5

\ddot{A}

IP Buffer: - + - + - + - + - + - + - + AGN194204

Stimulation: 10⁻⁶ 10⁻⁷ 10⁻⁸ 10⁻⁹ 10⁻¹⁰ 10⁻¹¹ (M) AGN194204 (10 min)



Flag-RXR α

[illegible]

FIGURE 6

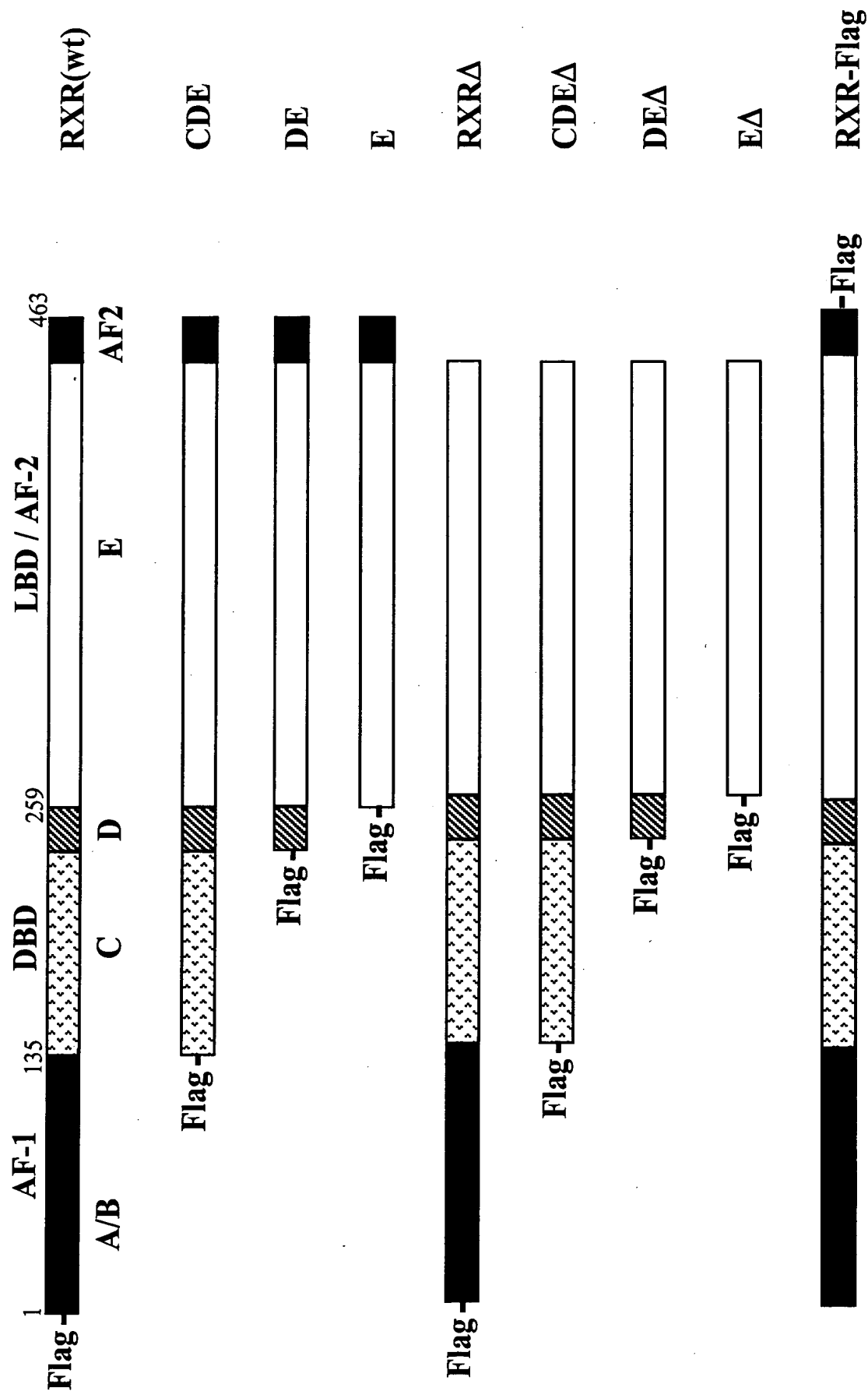
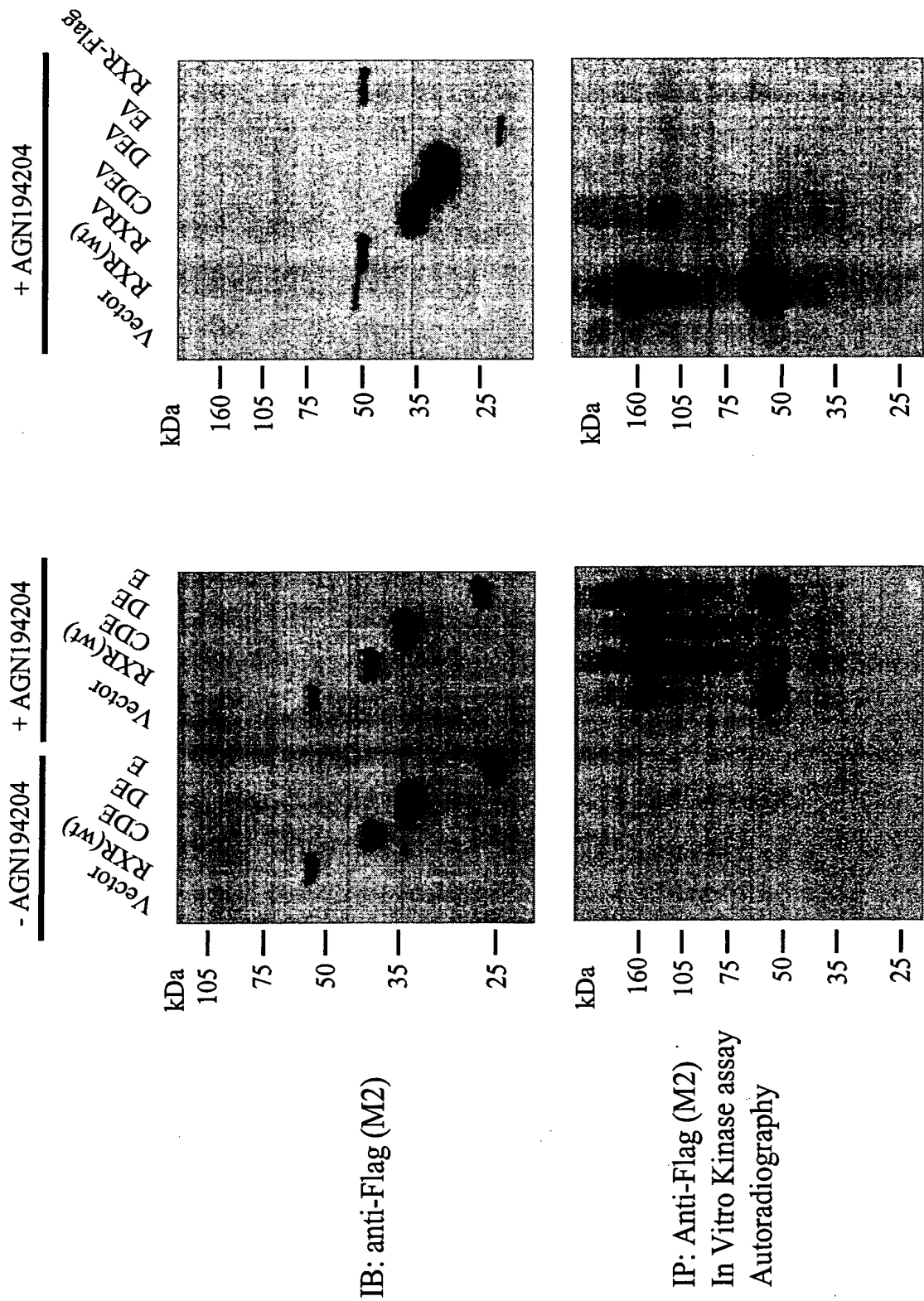


FIGURE 7

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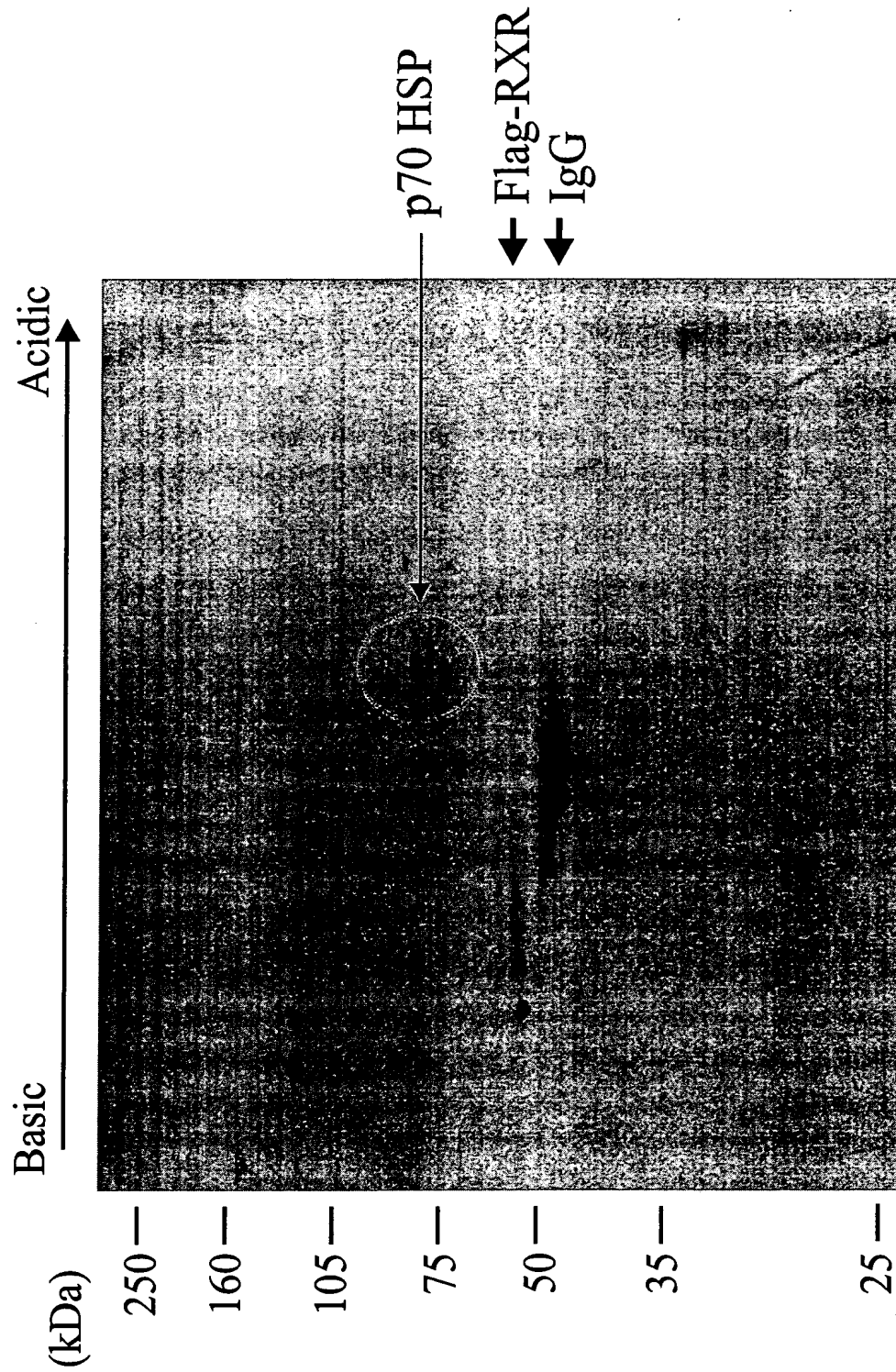


FIGURE 9

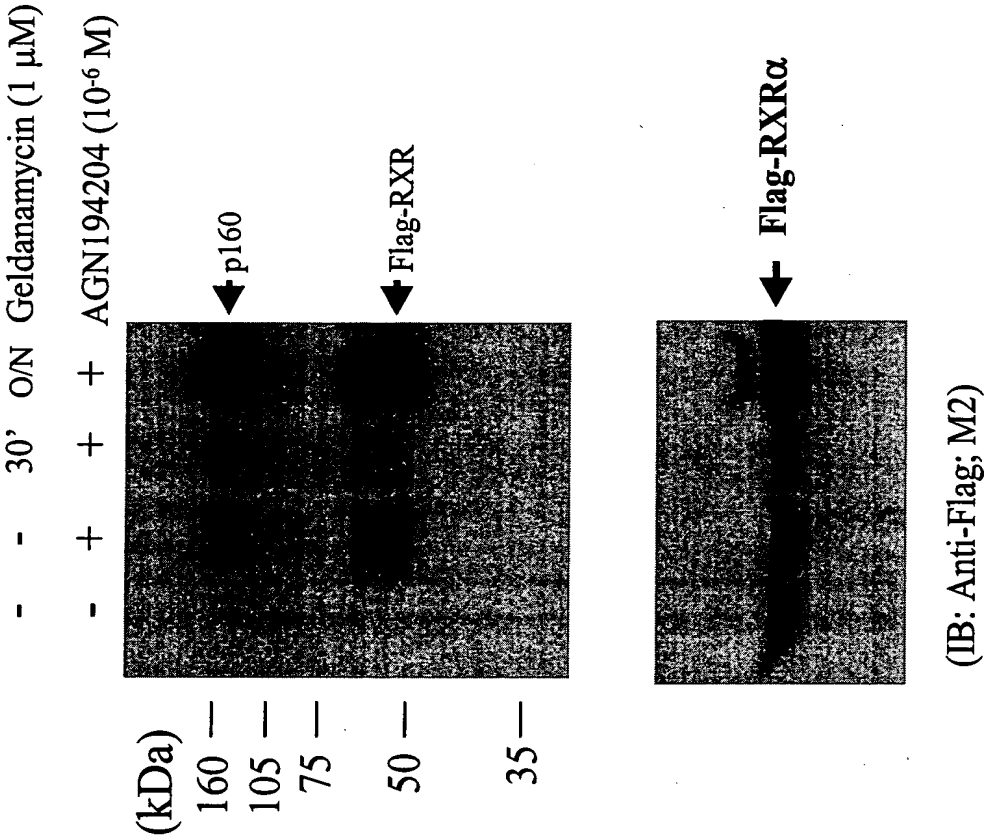


FIGURE 10

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A:

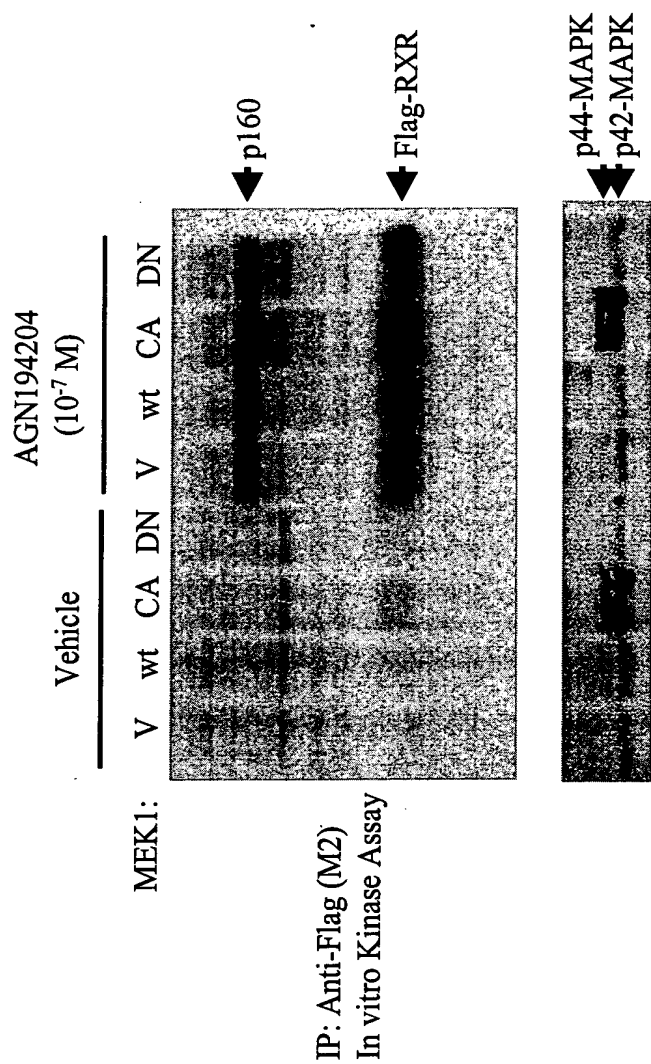


FIGURE 11

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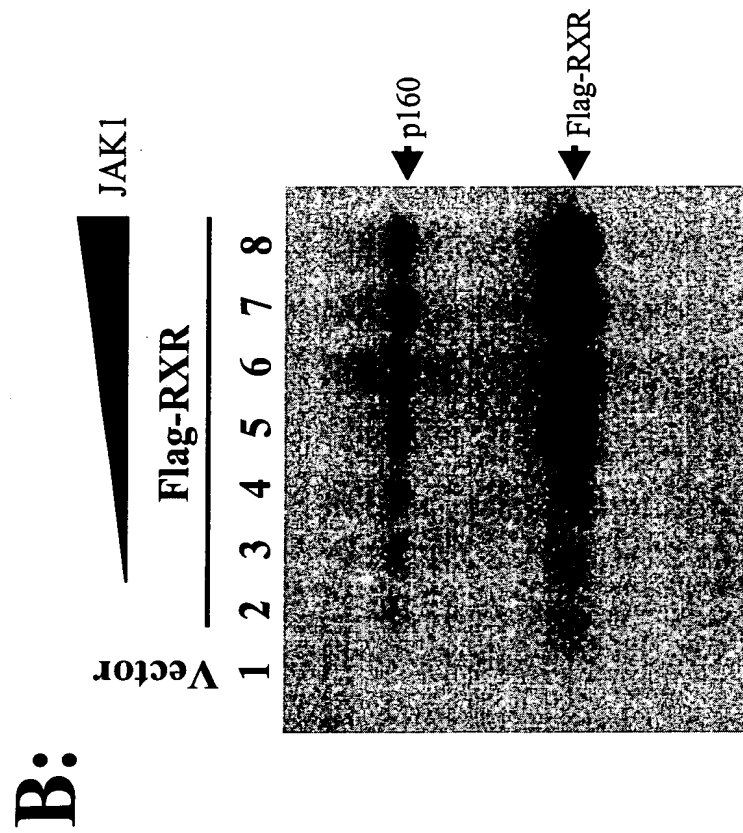
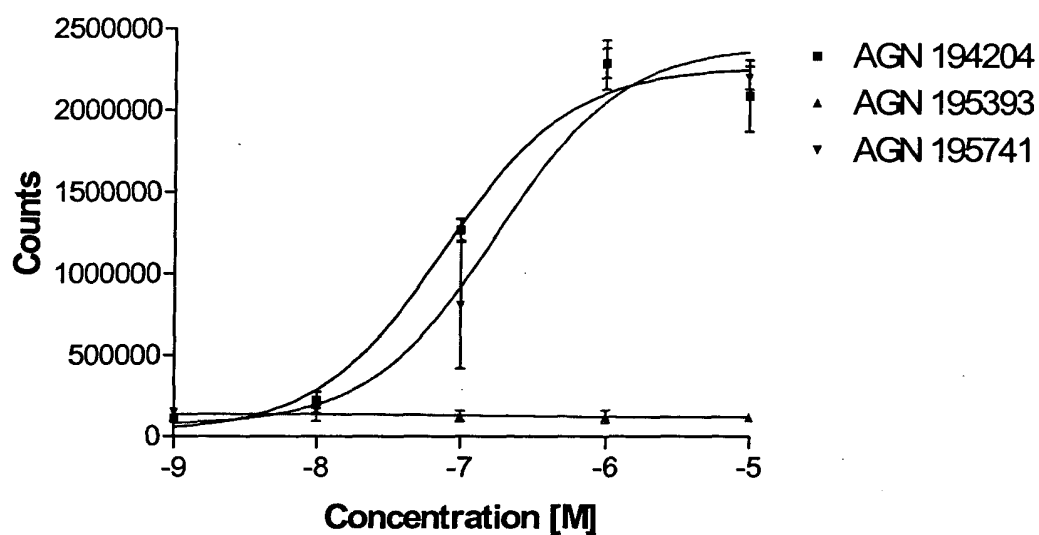
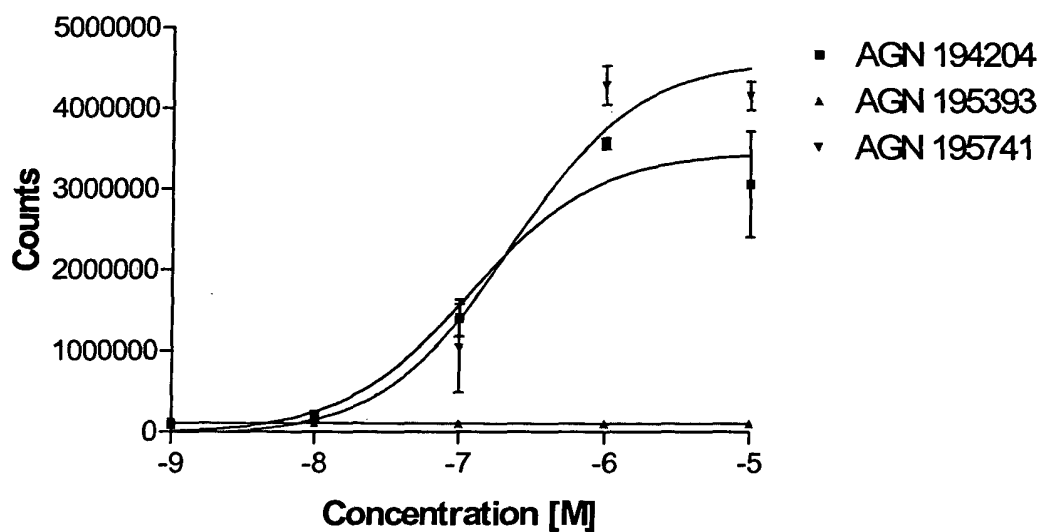


FIGURE 11 CONTINUED

A

	AGN 194204	AGN 195393	AGN 195741
EC50	7.7150e-008		1.7540e-007

B

	AGN 194204	AGN 195393	AGN 195741
EC50	1.2010e-007		2.2510e-007

FIGURE 12

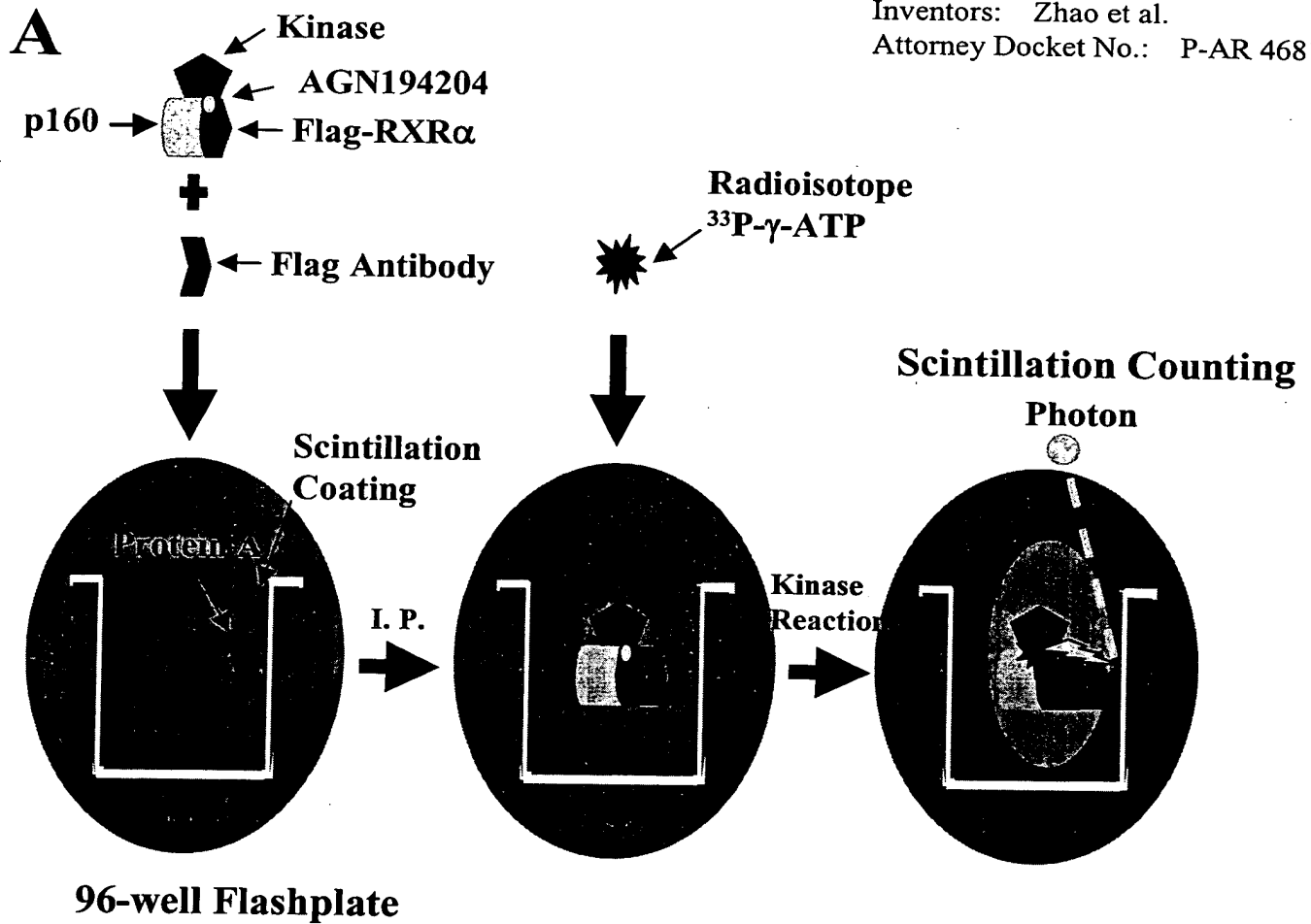
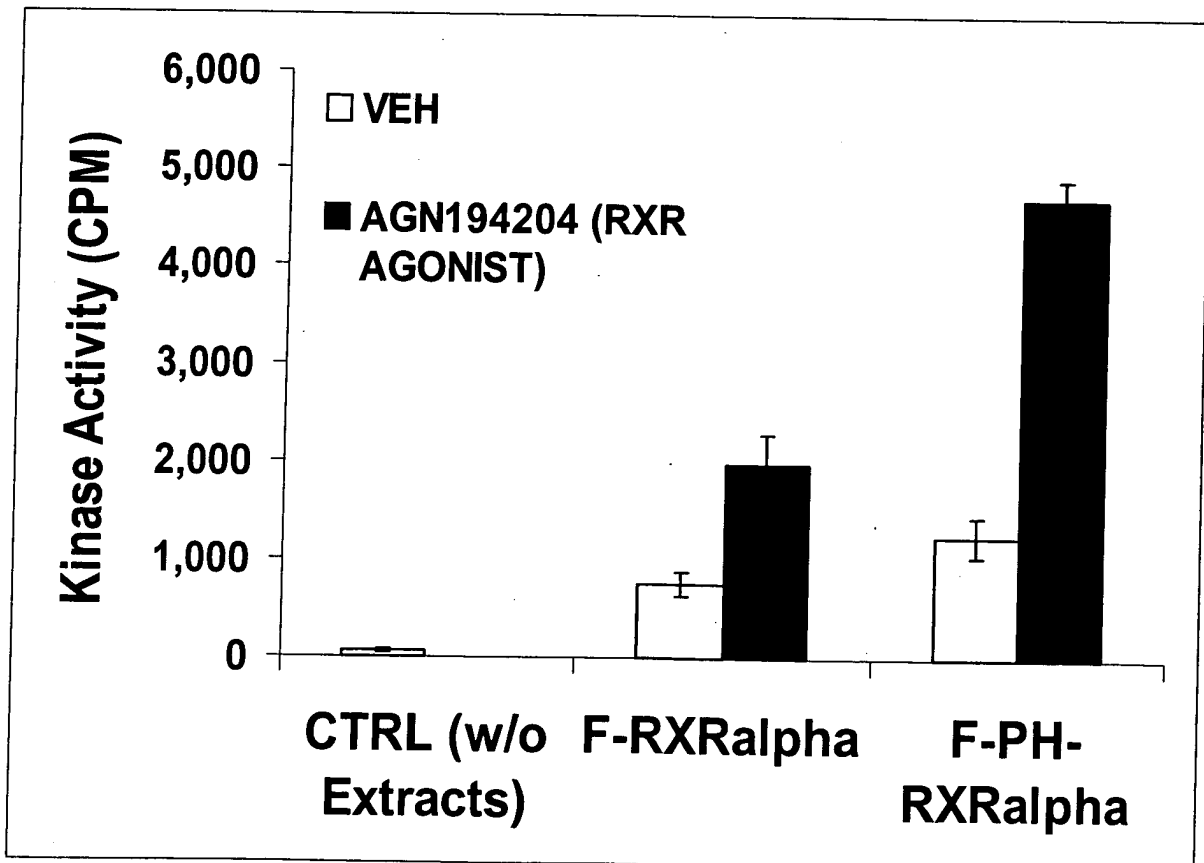
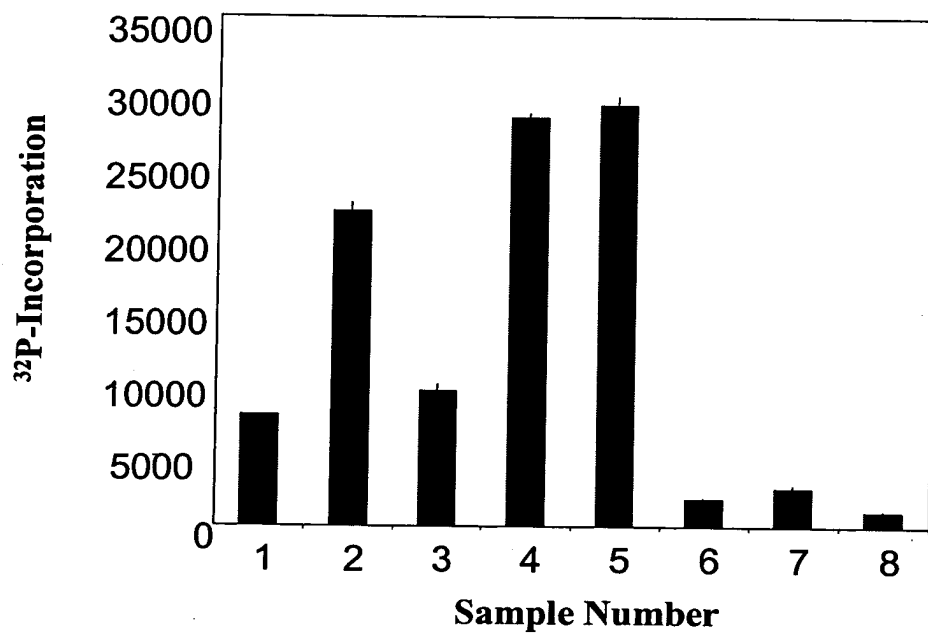
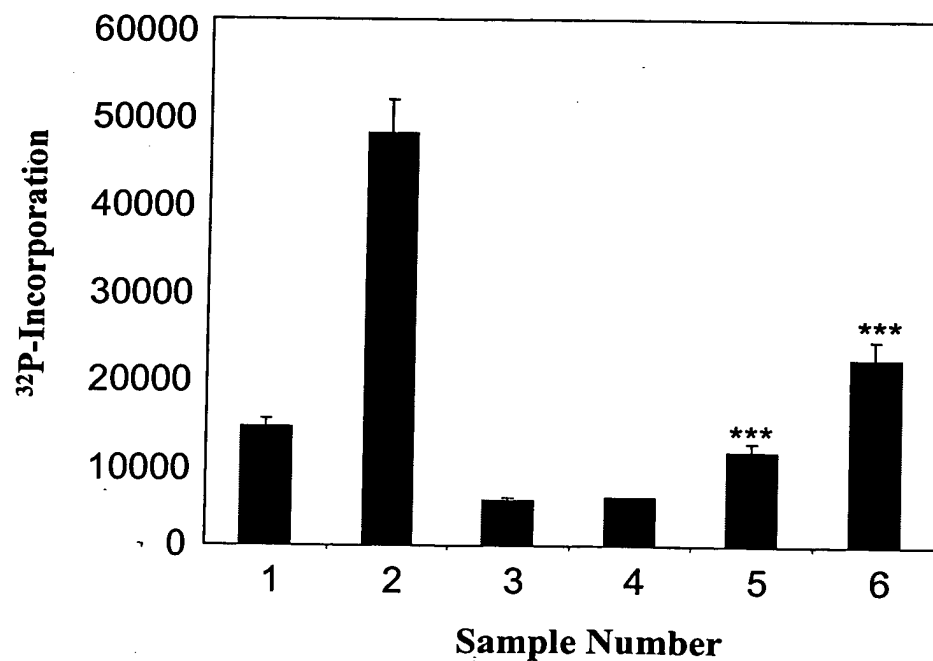
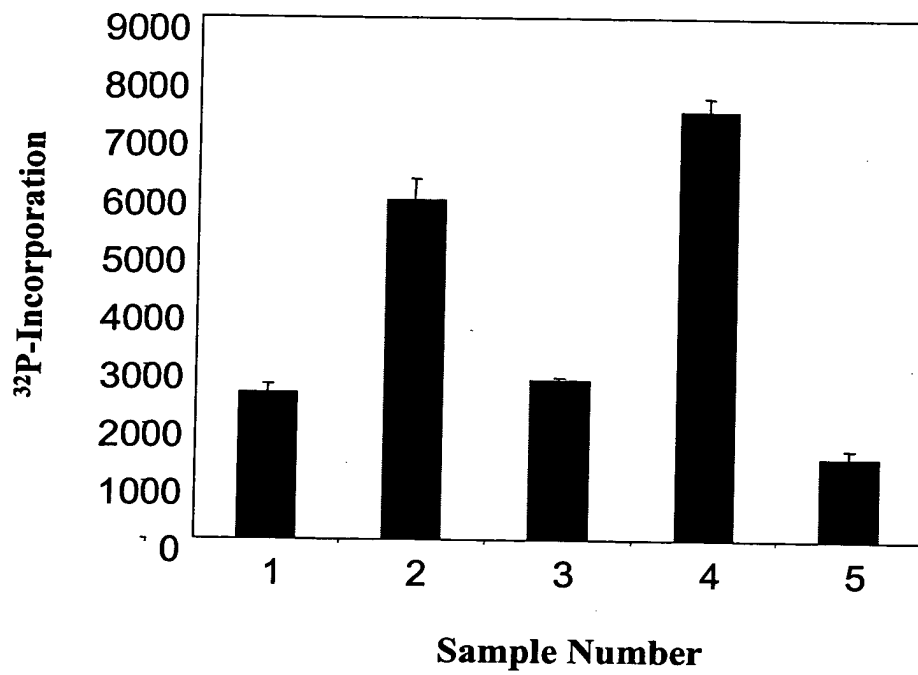
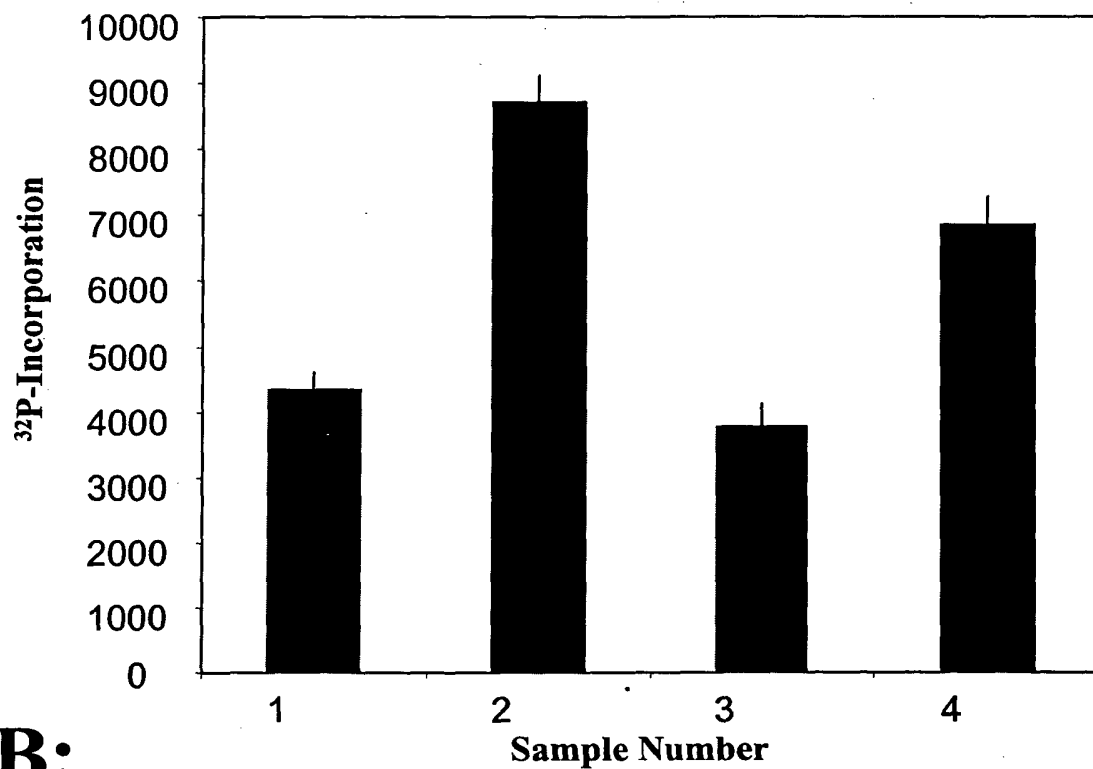
**B**

FIGURE 13

A:**B:****FIGURE 14**

**FIGURE 14 CONTINUED**

A:



B:

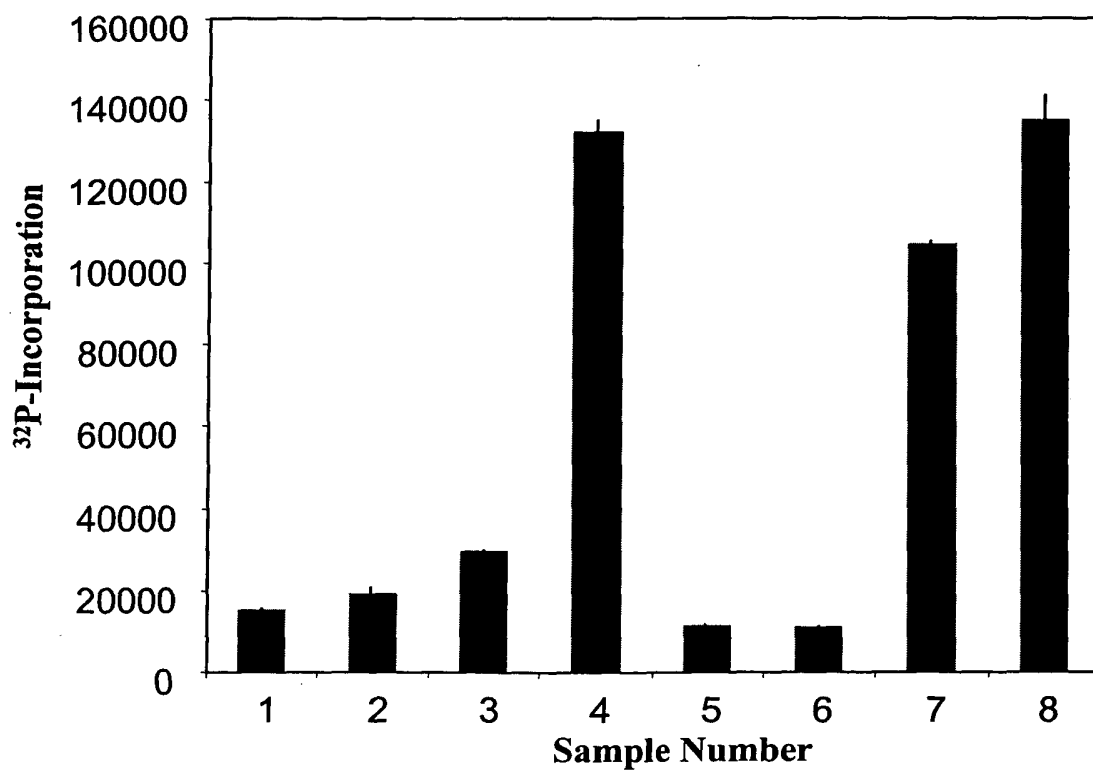


FIGURE 15

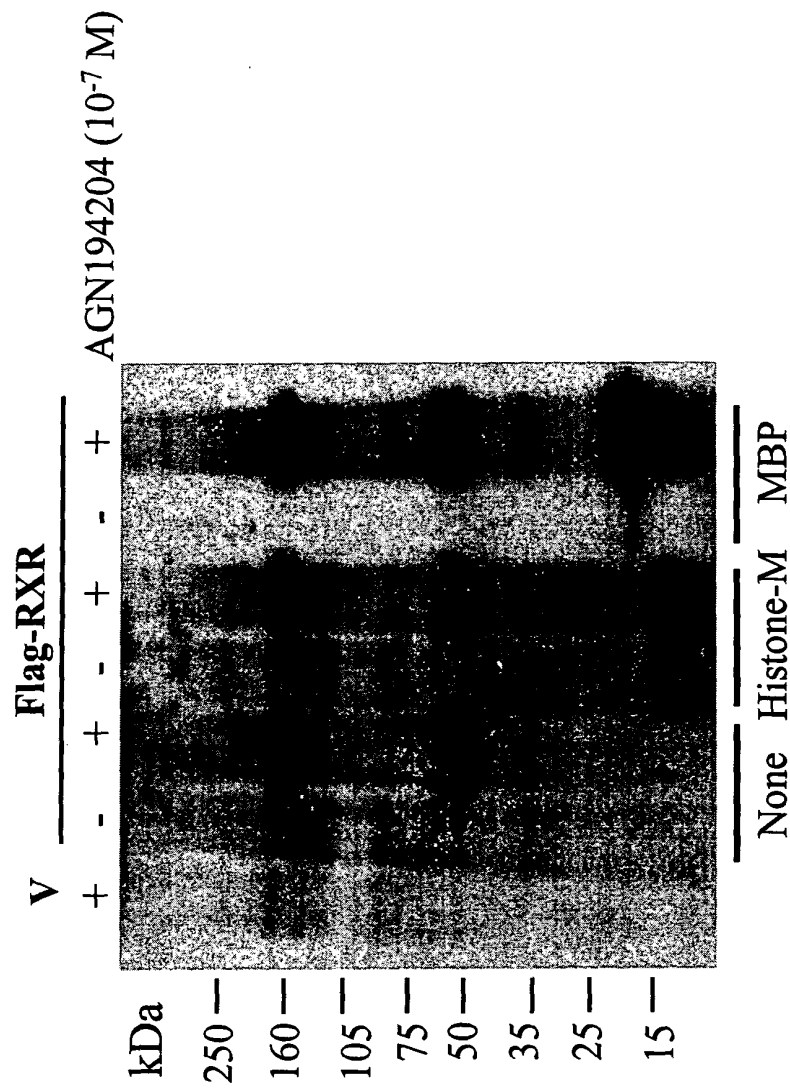
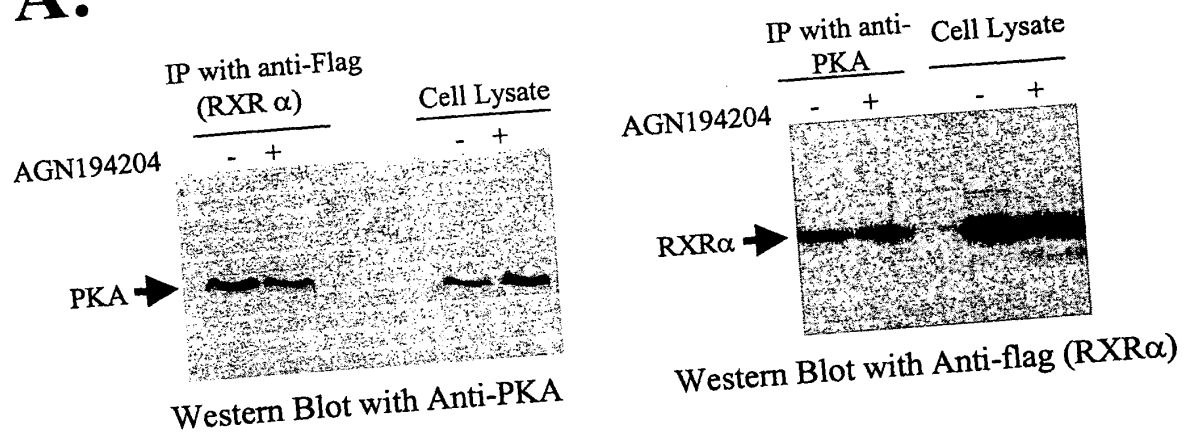


FIGURE 16

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A:



B:

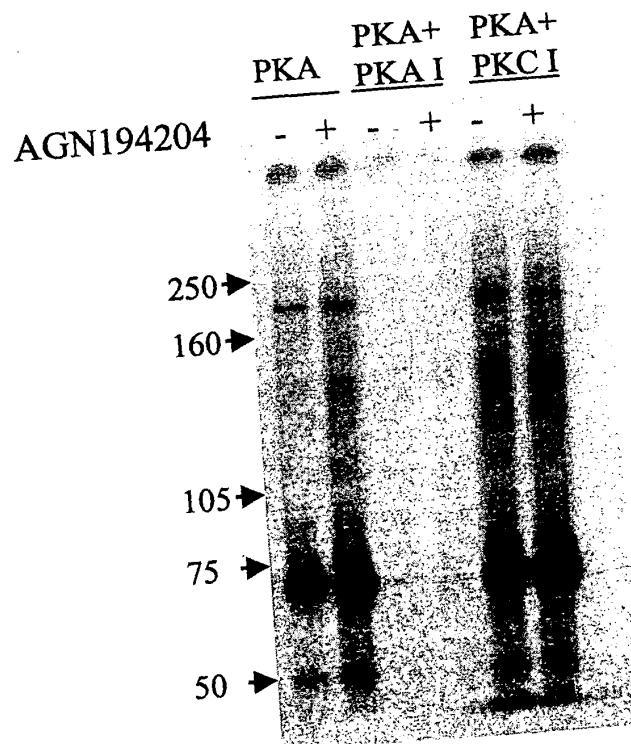


FIGURE 17

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Plec-N	GSTL	ISPCQD	FGKR	MEVFKIT	TTKQQ	DHFFQAAF	LEERDAMVRD	INKAIKICIEG
Plec-C	GCVVT	SVESN	SNGR	K	SEENLFEII	TADEV	HYFLOAAIT	IQMASRTGK*
RasGAP	VCSVY	VVHDS	LFGR	PNCFOIV	VQHFS	...EE	HYIFYFAGET	PECAEDMYKG	LOAFCNLRKS
Akt	NFSVA	ECQLM	KTER	PRPNTFVIR	CLQWT	...T	VIERTFHVDS	PDEREEMRA	IQMVANSLKQ
Spectrin	EAVCE	VALDY	KKKK	HVFKL	R	LNDGN	EYTFQAKD	ISSATSSDKI
BARK	EIQS	VEETQI	KER	KCLLK	IRGGK	QFIIQDCDS	DPELVQMKKE	LRCAVREAAQ
Tiam1-PH1	ENST	IVQVPE	HPKK	DFVFCLS	NSLGD	AFLFQTTSQ	TELENW	ITAI
Grb7	ESNVY	VVTQG	RKLY	G	MPTDFGFCVK	PNKLR	NGH	KGLHIFCSED	EDSRTCMYAA
Dynamamin	NLKL	RDVEKG	FMSS	KHIFALF	NTEQRNVYKD	YROLELACET	QEEVDSMKAS	FLRAGVYPER	
Irs1	SCFN	INKRAD	SKNK	HLVALY	TRDEH	FALIAACS	EAEQDSMYQA	LLQLHNRKA
RasGRF-PH1	EGS	ICKRMP	PKRG	TSSKES	DKQHYFTW	FSNDS	QKSLELRIDD	SKDCDENVAA	IARASYKILA
RasGRF-PH2	DCT	LLDPEN	MDDD	..GKGQ	EVDHLDKIM	VEPKD	...SP	PFTVILVASS	ROEKAAMTSD	IIQQVDNIR
Db1	EVG	ITEYVKG	DN	RKFEIM	YGEKE	EVYIVQASN	VDKMTMDKE	IRNILLKQOE
Vav	SFQV	RDDSSG	ERDN	KKWSHMLLI	EDQGA	QGVELFFKI	RELKKKIMEQ	FEMATISNIYP
Sos	KVQ	INDKDDT	NEYK	HAFETI	LKDEN	SVTFSAKS	AEEKNNIMAA	LISLQYRSTL
Bcr	QIKSD	IQREK	RANK	<19>LL	MSPSMAFRMH	SRNGK	SVTFLISS	DYERAEMREN	IREQQKKCFR
Btk	KITC	VETVVP	EKNP	<20>II	ERFPYPFQW	YDEGP	LYMFSPT	EEQKRMIHQ	LKNVIRYNSD
PLC-delta	DIQE	VRMGHR	TEGLE	KFARD	IPEDRCFSIV	FKDQRN	TLDLIAPS	PRDAQHMVQG	LRKIITHSGS
PLC-gamma	EIKE	IRPGKT	SRDF	<7>AFR	PDQSHCFVIL	YGMF	...RL	KTLSLOATSE	DEVNMIIKG	ITWLMEDTLQ

FIG.18A

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Plec-N	..MEPKRIRE	GYLVKKGVS.FNT	WKPMWVILEDGI	EFYKKK.SD	NSPKGMIPLK
Plec-C	EEFRGVIKQ	GCLLKQHR.RKN	WKVRKFIIREDPAYL	HYQPA.GA	EDPLGATHLR
RasGAP	DAFYKNIVKK	GYLLKKGKG.IKR	WKNLVFIILEGSDAQL	IYFESIKRA	TKPKGLICLS
Akt	.MNEVSVIKE	GMLHKRGEY.IKT	WRP YFQLKSDGSFI	GKERP.E	APDQTLPPIN
Spectrin	TQETPSAQME	GFLNRKHEWE	AHNKKASSRS	WHNVYCVINNQEM	GFYKDA.KTAASGIP	YHSEVPVSLK
BARK	YALGKDCIMH	GYMSKMGNP.FLTQ	WQRYFYIFPNRL	EMRGE.EAPQSLLTME	
Tiam1-PH1	VRKAGALAVK	NFLVHKKNK.	.KVESATPRK	WKHYMWSLKGCTL	FFYETD.GRSRIDH	NSVPKHAVW
Grb7	LNAGSFPEIQ	GFLQLRGSG.	...RGSGRKL	WKRFCCFLRRSGL	YYSTKG.TSKDP	RHLQYVADV
Dynammin	NQDEILVIRK	GMLTINNIG.	...IMGG	SKEYMFVILTAENL	SWYKDD.EE	KEKRYMISMD
Irs1	TDGFSDVRKV	GYLRKPKS..	MHKRFVILRA	ASEAGGPAL	EYENE.KKWRHKS	SAPKRSIPLE
RasGRF-PH1	LLAQRDGTRK	GYLSKRSSD.NPK	WQTRWFAILLQNLL	FYQESD.S	SSRPSGIYLL
RasGRF-PH2	LDTSQTFVRQ	GSLIQVPMS	<10>GSLSLKK	EGEQCFILFSKHL	IICTRG.SGSKLH	LTKNGVLSLI
Db1	IGHKKGATKM	KDLARFK...	PMQRHIFLYEKAT	VECKRR.	VE	SGEGSDRYP
Vav	LANYGRPKID	GELKITSVE.RRS	KTDRAVFLDKAL	LICKRR.GDS	YDLKASVNLH
Sos	GQCCNEFTME	GILTRVGA..	KHERHIFLFDGL	MICCKSNHGQ	PRLPGASNAE	YRLKEKFFMR
Bcr	VFLFTDILLC	TKLKQSGG.KTQQ	YDTCKVYPLTDL	SFQWDELEA	VPNIPLVPDE	ELDALKIKLS
Btk	..MAAVILES	IFLKRQQK.	...KKTSPN	FKRQELLTVHKL	SWVEYD.FERGR	GSKKGSIDME
PLC-delta	DDPDLOALIK	GSQLLKVK.SSW	RRRFYKIQEDCK	TINGES.RKVMR	SPEQLFSLE
PLC-gamma	VLHLCRSLLEV	GTMTLFYS.	...KKSQPE	RKTFQVKLETROL	TSRGA.	DKIEGALDIR

FIG.18B